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#### **Declarations under Rule 4.17:**

- as to applicant's entitlement to apply for and be granted a patent (Rule 4.17(ii))
- as to the applicant's entitlement to claim the priority of the earlier application (Rule 4.17(iii))

**Published:** 

- with international search report
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(54) Title: ALKOXY SUBSTITUTED IMIDAZOQUINOLINES

(57) Abstract: Imidazoquinoline compounds with an alkoxy substituent at the 6, 7, 8, or 9-position, pharmaceutical compositions containing the compounds, intermediates, methods of making, and methods of use of these compounds as immunomodulators, for inducing or inhibiting cytokine biosynthesis in animals and in the treatment of diseases including viral, and neoplastic, are disclosed.

63. The compound or salt of claim 55 or claim 62 wherein  $R_6$  is =0 or =S, a and b are each independently 1 to 3, and A' is selected from the group consisting of  $-CH_2$ -,  $-S(O)_2$ -, and -O-.

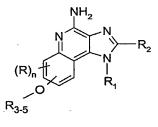
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64. The compound or salt of claim 55, claim 62, or claim 63 wherein  $Z_a$  is methylene,  $R_6$  is =0, a is 1 or 2, b is 2, and A' is -CH<sub>2</sub>-.

65. The compound or salt of claim 55, claim 62, or claim 63 wherein  $Z_a$  is methylene, 10 R<sub>6</sub> is =O, a and b are each 2, and A' is -O-.

66. The compound or salt of claim 55 wherein  $Z_a$  is a bond or alkylene.

67. A compound of the formula (VII):



VΠ

wherein:

 $R_{3-5}$  is selected from the group consisting of

-Z-N(R<sub>8</sub>)-C(R<sub>6</sub>)-O-R<sub>4</sub>,  

$$-Z - \left( \begin{array}{c} N - C(R_6) - O - R_4 \\ R_{10} \end{array} \right)$$
, and

 $-Z-N(R_8)-C(R_6)-C(R_6)-R_4;$ 

Z is selected from the group consisting of alkylene, alkenylene, and alkynylene, wherein alkylene, alkenylene, and alkynylene can be optionally interrupted with one or more -O- groups;

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R is selected from the group consisting of alkyl, alkoxy, hydroxy, halogen, and trifluoromethyl;

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# PATENT COOPERATION TREATY

# PCT

### INTERNATIONAL PRELIMINARY REPORT ON PATENTABILITY (Chapter I of the Patent Cooperation Treaty)

(PCT Rule 44bis)

Applicant's or agent's file reference 58915WO006	FOR FURTHER ACTION	See item 4 below	
International application No. PCT/US2004/032616	International filing date (day/month/year) 01 October 2004 (01.10.2004)	Priority date (day/month/year) 03 October 2003 (03.10.2003)	
International Patent Classification (8t See relevant information in Form I	h edition unless older edition indicated) PCT/ISA/237	······································	
Applicant 3M INNOVATIVE PROPERTIES (	COMPANY		

1.	This international preliminary report on patentability (Chapter I) is issued by the International Bureau on behalf of the International Searching Authority under Rule 44 <i>bis</i> . 1(a).		
2.	This REPORT consists of a total of 7 sheets, including this cover sheet.		
	In the attached sheets, any refer to the international preliminary	rence to the written opinion of the International Searching Authority should be read as a reference report on patentability (Chapter I) instead.	
3.	This report contains indications	relating to the following items:	
	Box No. I	Basis of the report	
	Box No. II	Priority	
	Box No. III	Non-establishment of opinion with regard to novelty, inventive step and industrial applicability	
	Box No. IV	Lack of unity of invention	
	Box No. V	Reasoned statement under Article 35(2) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement	
	Box No. VI	Certain documents cited	
	Box No. VII	Certain defects in the international application	
	Box No. VIII	Certain observations on the international application	
4.	The International Bureau will cont, except where the applicant date (Rule 44 <i>bis</i> .2).	ommunicate this report to designated Offices in accordance with Rules 44 <i>bis</i> .3(c) and 93 <i>bis</i> .1 but makes an express request under Article 23(2), before the expiration of 30 months from the priority	

<b>-</b>	Date of issuance of this report 03 April 2006 (03.04.2006)
The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland	Authorized officer Athina Nickitas-Etienne
Facsimile No. +41 22 740 14 35	Telephone No. +41 22 338 89 95

Form PCT/IB/373 (January 2004)

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# PATENT COOPERATION TREATY

From the INTERNATIONAL SEARCHING AUTH	IORITY		*** *		
INTERNATIONAL SEARCHING AUTHORITY To: DEAN A. ERSFELD OFFICE OF INTELLECTUAL PROPERTY COUNSEL			РСТ	REC'D 20 APR	2005
POST OFFICE BOX 33427 SAINT PAUL, MN 55133-3427	RIY COUNSEL	WR INTERNATIO	ITTEN OPINION ONAL SEARCHIN	WIPO OF THE IG AUTHORITY	PCT
			(PCT Rule 43bis.	1)	
		Date of mailing (day/month/year)	18 APR 2		7
Applicant's or agent's file reference		FOR FURTHER	ACTION		-
58915WO006			See paragraph 2 below		
International application No.	International filing date	(day/month/year)	Priority date (day/mo	nth/year)	
PCT/US04/32616 International Patent Classification (IPC)	01 October 2004 (01.10	.2004)	03 October 2003 (03.	10.2003)	4
IPC(7): A61K 31/4745, 31/496, 31/506, 514/293, 232.8, 253.03, 218, 214.03, 2 Applicant	, 31/5377, 31/541, 31/551 17.07, 273, 228.5; 546/82	; C07D 471/04, 401, 2; 544/126, 60, 361;	/14, 405/14; A61P 37/0 540/575, 585, 597	02 and US C1.:	
3M INOVATIVE PROPERTIES COMP	ANY				
1. This opinion contains indications rel	ating to the following iten	ns:			]
Box No. I Basis of the opinion					
Box No. II Priority					
Box No. III Non-establishment of opinion with regard to novelty, inventive step and industrial applicability					
Box No. IV Lack of unity of invention					
	tatement under Rule 43bis y; citations and explanatio			p or industrial	
Box No. VI Certain doc	uments cited				
Box No. VII Certain def	ects in the international ap	plication			
Box No. VIII Certain obs	ervations on the internatio	nal application			
2. FURTHER ACTION					
If a demand for international prelim International Preliminary Examinin Authority other than this one to be that written opinions of this Internation	g Authority ("IPEA") ex the IPEA and the chosen l	ccept that this does PEA has notified the	not apply where the International Bureau	applicant chooses an	
If this opinion is, as provided above IPEA a written reply together, wh mailing of Form PCT/ISA/220 or be	ere appropriate, with am fore the expiration of 22	endments, before th	e expiration of 3 mon	ths from the date of	
For further options, see Form PCT/	ISA/220.				
3. For further details, see notes to Form	n PCT/ISA/220.				
· · · · · · · · · · · · · · · · · · ·	Name and mailing address of the ISA/ US Authorized officer				1
Mail Stop PCT, Attn: ISA/US Commissioner for Patents		Evelyn Huang	7. Rober	to for	
P.O. Box 1450 Alexandria, Virginia 22313-1450		Telephone No. (5		V	
Facsimile No. (703) 305-3230 Form PCT/ISA/237 (cover sheet) (January	2004)	L	··· ··· ··· ··· ··· ··· ··· ··· ··· ··		1

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	WRITTEN OPINION OF THE	International application No.
·	INTERNATIONAL SEARCHING AUTHORITY	PCT/US04/32616
Box N	No. I Basis of this opinion	
1. With it wa	regard to the language, this opinion has been established on the basis of as filed, unless otherwise indicated under this item. This opinion has been established on the basis of a translation from the which is the language of a translation furnished for the aver	original language into the following language
	, which is the language of a translation furnished for the purp 23.1(b)).	
2. With claim	regard to any nucleotide and/or amino acid sequence disclosed in ned invention, this opinion has been established on the basis of:	the international application and necessary to the
a.	type of material	
	a sequence listing	
	table(s) related to the sequence listing	
Ъ.	format of material	
	in written format	
	in computer readable form	
c.	time of filing/furnishing	
	contained in international application as filed.	
l	filed together with the international application in computer reade	able form.
	furnished subsequently to this Authority for the purposes of search	h.
3. 🛄	In addition, in the case that more than one version or copy of a sequirated or furnished, the required statements that the information in the su the application as filed or does not go beyond the application as filed.	ubsequent or additional copies is identical to that in
4. Additi	ional comments:	

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Form PCT/ISA/237(Box No. I) (January 2004)

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	WRITTEN OPINION OF THE	International application No.
	INTERNATIONAL SEARCHING AUTHORITY	PCT/US04/32616
Box N	o. III Non-establishment of opinion with regard to novelty, in	wentive step and industrial applicability
1. The	questions whether the claimed invention appears to be novel, to involve a strially applicable have not been examined in respect of:	
	the entire international application	
X	claims Nos. 5-10.14.21.25.34.38.41.64.65 and 76-90	
becar	use:	
	the said international application, or the said claim Nos rel require an international preliminary examination (specify):	late to the following subject matter which does no
$\boxtimes$	the description, claims or drawings ( <i>indicate particular elements below</i> ) and <u>76-90</u> are so unclear that no meaningful opinion could be formed (s They are improperly multiply dependent claims. PCT Rule 6.4	specify):
-		
	the claims, or said claims Nos are so inadequately supported	d by the description that no meaningful opinion
	could be formed.	
	no international search report has been established for said claims Nos. the nucleotide and/or amino acid sequence listing does not comply with Administrative Instructions in that:	ith the standard provided for in Annex C of th
	the written form has not been furnished	
	the computer readable form does not comply with the has not been furnished does not comply with the does not comply with	
	the tables related to the nucleotide and/or amino acid sequence listing, if	in computer readable form only, do not comply
	with the technical requirements provided for in Annex C-bis of the Adm	inistrative Instructions.

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WRITTEN OPINION OF THE	International application No.
INTERNATIONAL SEARCHING AUTHORITY	PCT/US04/32616
Box No. IV Lack of unity of invention	
<ol> <li>In response to the invitation (Form PCT/ISA/206) to pay addition paid additional fees</li> <li>paid additional fees under protest</li> <li>not paid additional fees</li> </ol>	al fees the applicant has:
<ol> <li>This Authority found that the requirement of unity of invention is to pay additional fees.</li> </ol>	not complied with and chose not to invite the applicant
<ol> <li>This Authority considers that the requirement of unity of invention in ac</li> <li>complied with</li> </ol>	cordance with Rule 13.1, 13.2 and 13.3 is
not complied with for the following reasons:	
See the lack of unity section of the International Search Report(Form PC	CT/ISA/210)
`	
<ol> <li>Consequently, this opinion has been established in respect of the following all parts.</li> </ol>	g parts of the international application:
the parts relating to claims Nos.	
orm PCT/ISA/237 (Box No. IV) (January 2004)	

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WRITTEN OPINION OF THE International application No. INTERNATIONAL SEARCHING AUTHORITY	
Box No. V Reasoned statement under Rule 43 bis.1(a)(i) with regard to novelty, inventive step or industria applicability; citations and explanations supporting such statement	əl
1. Statement	
Novelty (N) Claims Please See Continuation Sheet	YES
Claims Please See Continuation Sheet	_NO
Inventive step (IS) Claims Please See Continuation Sheet	_YES
Claims Please See Continuation Sheet	_NO
Industrial applicability (IA) Claims Please See Continuation Sheet	YES
Claims Please See Continuation Sheet	NO

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2. Citations and explanations:

Claims 1-4, 11-13, 15-20, 22-24, 26-33, 35-37, 39, 40, 42-63, and 66-75 meet the criteria set out in PCT Article 33(2)-(3), because the closest prior art, US 6,200,592 B1, only teaches alkoxy, and does not teach or fairly suggest the instant -O-R<sub>3</sub> substituent on the imidazoquinoline.

Claims 91-93 meet the criteria set out in PCT Article 33(2)-(3), because the closest prior art, US 4,698,348 A, only teaches methoxy, and does not teach or fairly suggest the instant -O-R<sub>3</sub> substituent on the imidazoquinoline.

Claims 1-4, 11-13, 15-20, 22-24, 26-33, 35-37, 39, 40, 42-63, 66-75, and 91-93 meet the criteria set out in PCT Article 33(4), and thus find industrial applicability because the subject matter claimed can be made or used in the pharmaceutical industry as immunomodulating agent in treatment of diseases.

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Form PCT/ISA/237 (Box No. V) (January 2004)

# WRITTEN OPINION OF THE INTERNATIONAL SEARCHING AUTHORITY

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International application No. PCT/US04/32616

Supplemental Box In case the space in any of the preceding boxes is not sufficient.
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V.1. Reasoned Statements: The opinion as to Novelty was positive (Yes) with respect to claims 1-4, 11-13, 15-20, 22-24, 26-33, 35-37, 39, 40, 42-63, 66-75, 91-
93
The opinion as to Novelty was negative (No) with respect to claims NONE The opinion as to Inventive Step was positive (Yes) with respect to claims 1-4, 11-13, 15-20, 22-24, 26-33, 35-37, 39-40, 42-63, 66-
75, 91-93 The opinion as to Inventive Step was negative (NO) with respect to claims NONE
The opinion as to Industrial Applicability was positive (YES) with respect to claims 1-4, 11-13, 15-20, 22-24, 26-33, 35-37, 39, 40, 42-63, 66-75, 91-93
The opinion as to Industrial Applicability was negative(NO) with respect to claims NONE

Form PCT/ISA/237 (Supplemental Box) (January 2004)